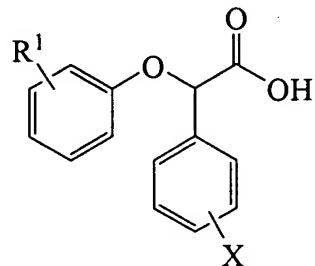


Amendments to the Claims:

This listing of claims will replace all prior versions, and listings of claims in the application:

Listing of Claims:

1. (Cancel) A method for producing an enantiomerically enriched α -(phenoxy)phenylacetic acid compound of the formula:



wherein

R^1 is alkyl or haloalkyl, and

X is halide;

from an enantiomeric mixture of the α -(phenoxy)phenylacetic acid compound comprising a first and a second enantiomers, said method comprising:

(a) producing a solution comprising a solid enantiomerically enriched acid-base salt of the first enantiomer by contacting the enantiomeric mixture of the α -(phenoxy)phenylacetic acid compound with less than 0.5 molar equivalents of an enantiomerically enriched chiral amine compound under conditions sufficient to produce the ratio of the amount of free first enantiomer to the amount of the free second enantiomer in the solution is about 1 to 3; and

(b) separating the solid acid-base salt of the first enantiomer from the solution at a temperature where the concentration of an acid-base salt of the second enantiomer of the α -(phenoxy)phenylacetic acid compound is near or below its saturation point.

2. (Cancel) The method of Claim 1, wherein said step (a) of producing the solution comprising the solid enantiomerically enriched acid-base salt of the first enantiomer comprises:

- (i) heating the solution to a temperature above the nucleation temperature of the first enantiomer; and
- (ii) lowering the solution temperature to a temperature at or below the nucleation temperature of the first enantiomer to produce the solid acid-base salt of the first enantiomer.

3. (Cancel) The method of Claim 2, wherein said step (b) of separating the solid acid-base salt of the first enantiomer is conducted at a temperature near or above a saturation temperature of an acid-base salt of the second enantiomer.

4. (Cancel) The method of Claim 1 further comprising recovering the chiral amine compound by removing the chiral amine compound from the separated solid acid-base salt of the first enantiomer.

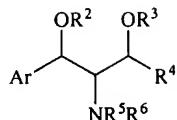
5. (Cancel) The method of Claim 4, wherein the enantiomerically enriched chiral amine compound used in producing the acid-base salt of said step (a) comprises the recovered chiral amine compound.

6. (Cancel) The method of Claim 1 further comprising racemizing at least a portion of the second enantiomer in the separated solution by contacting the second enantiomer with a base.

7. (Cancel) The method of Claim 6, wherein the enantiomeric mixture of the α -(phenoxy)phenylacetic acid compound used in said step (a) comprises a racemized α -(phenoxy)phenylacetic acid compound.

8. (Cancel) The method of Claim 1, wherein the chiral amine compound is of the formula:

Amdt. dated December 22, 2005

Response to Restriction Requirement and Preliminary
Amendment

wherein

each of R² and R³ is independently hydrogen or alkyl; or R² and R³ together with atoms to which they are attached to form a heterocyclic ring moiety;

R⁴ is hydrogen or alkyl;

each of R⁵ and R⁶ is independently hydrogen or alkyl, or one of R⁵ or R⁶ is an amine protecting group; and

Ar is aryl.

9. (Cancel) A method for enantiomerically enriching (-)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid from an enantiomeric mixture of 4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid, said method comprising:

(a) producing a solution comprising an enantiomerically enriched acid-base salt of (-)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid by contacting the enantiomeric mixture of 4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid with less than 0.5 molar equivalent of an enantiomerically enriched (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol in about 4 grams of an alcoholic solvent per gram of (-)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid;

(b) separating the enantiomerically enriched acid-base salt from the solution which is enriched with (+)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid; and

(c) removing (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol from the acid-base salt to produce enantiomerically enriched (-)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid.

10. (Cancel) The method of Claim 9, wherein the alcoholic solvent is isopropanol.

11. (Cancel) The method of Claim 10, wherein about 0.47 molar equivalent or less of (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol is used to form the acid-base salt.

Amdt. dated December 22, 2005

Response to Restriction Requirement and Preliminary
Amendment

12. (Cancel) The method of Claim 11, wherein said step (a) of producing a solution comprising an enantiomerically enriched acid-base salt of (-)-4-chloro- α -(3-trifluoromethyl-phenoxy)phenylacetic acid comprises heating the solution mixture to a temperature at or above a nucleation temperature of the (-)-acid-base salt.

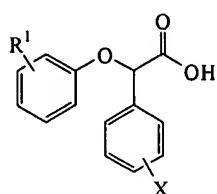
13. (Cancel) The method of Claim 12, wherein said step (b) of separating the enantiomerically enriched acid-base salt is performed at a temperature near or above a saturation temperature of an acid-base salt of the (+)-enantiomer.

14. (Cancel) The method of Claim 10, wherein the enantiomerically enriched (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol comprises at least a portion of (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol that is removed from the acid-base salt of said step (c).

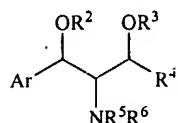
15. (Cancel) The method of Claim 10 further comprising racemizing at least a portion of (+)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid obtained in said step (b).

16. (Cancel) The method of Claim 15, wherein the enantiomeric mixture of 4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid comprises at least a portion of (+)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid that is racemized.

17. (Cancel) An acid-base salt derived from a α -(phenoxy)phenylacetic acid compound of the formula:



and a chiral amine compound of the formula:



wherein

Amdt. dated December 22, 2005

Response to Restriction Requirement and Preliminary
AmendmentR¹ is alkyl or haloalkyl;

X is halide;

each of R² and R³ is independently hydrogen or alkyl; or R² and R³ together with atoms to which they are attached to form a heterocyclic ring moiety;R⁴ is hydrogen or alkyl;each of R⁵ and R⁶ is independently hydrogen or alkyl, or one of R⁵ or R⁶ is an amine protecting group; and

Ar is aryl.

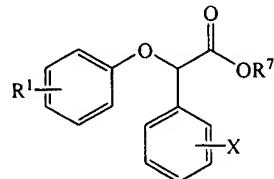
18. (Cancel) The acid-base salt of Claim 17, wherein the α -(phenoxy)phenylacetic acid compound and the chiral amine compound are enantiomerically enriched.

19. (Cancel) The acid-base salt of Claim 18, wherein the α -(phenoxy)phenylacetic acid compound is (-)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid.

20. (Cancel) The acid-base salt of Claim 18, wherein the chiral amine compound is (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol.

21. (Cancel) An enantiomerically enriched (-)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid having an enantiomeric excess of at least about 95%.

22. (Currently amended) A process for enantioselectively producing a α -(phenoxy)phenylacetate compound of the formula:

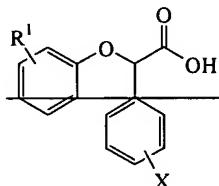


said method comprising:

Amdt. dated December 22, 2005

Response to Restriction Requirement and Preliminary
Amendment

(a) ~~producing a racemic mixture of a α -(phenoxy)phenylacetic acid of the formula:~~



(b)—resolving the racemic mixture of the α -(phenoxy)phenylacetic acid using less than 0.5 molar equivalent of an enantiomerically enriched chiral amine compound to produce an enantiomerically enriched α -(phenoxy)phenylacetic acid;

(be) producing an enantiomerically enriched activated α -(phenoxy)phenylacetic acid by contacting the enantiomerically enriched α -(phenoxy)phenylacetic acid with a carboxylic acid activating reagent; and

(cd) contacting the enantiomerically enriched activated α -(phenoxy)phenylacetic acid with a compound of the formula $(R^7-O)_wM$ to produce the α -(phenoxy)phenylacetate compound,

wherein

R^1 is alkyl or haloalkyl;

X is halide;

R^7 is heteroalkyl;

M is hydrogen or a metal; and

the subscript w is the oxidation state of M.

23. (Original) The method of Claim 22, wherein the α -(phenoxy)phenylacetate compound is (-)-halofenate.

24. (New) The method of Claim 22, wherein said step (a) resolving the racemic mixture of the α -(phenoxy)phenylacetic acid comprises:

(a) producing a solution comprising a solid enantiomerically enriched acid-base salt of a first enantiomer by contacting the enantiomeric mixture of the α -(phenoxy)phenylacetic acid compound with less than 0.5 molar equivalents of an

Amdt. dated December 22, 2005

Response to Restriction Requirement and Preliminary
Amendment

enantiomerically enriched chiral amine compound under conditions sufficient to produce the ratio of the amount of free first enantiomer to the amount of the free second enantiomer in the solution is about 1 to 3; and

(b) separating the solid acid-base salt of the first enantiomer from the solution at a temperature where the concentration of an acid-base salt of the second enantiomer of the α -(phenoxy)phenylacetic acid compound is near or below its saturation point.

25. (New) The method of Claim 24, wherein said step (a) of producing the solution comprising the solid enantiomerically enriched acid-base salt of the first enantiomer comprises:

(i) heating the solution to a temperature above the nucleation temperature of a first enantiomer; and

(ii) lowering the solution temperature to a temperature at or below the nucleation temperature of the first enantiomer to an enantiomerically enriched α -(phenoxy)phenylacetic acid.

26. (New) The method of Claim 24, wherein said step (b) of separating the solid acid-base salt of the first enantiomer is conducted at a temperature near or above a saturation temperature of an acid-base salt of the second enantiomer.

27. (New) The method of Claim 22 further comprising recovering the chiral amine compound by removing the chiral amine compound from the separated solid acid-base salt of the first enantiomer.

28. (New) The method of Claim 27, wherein the enantiomerically enriched chiral amine compound used in producing the acid-base salt of said step (a) comprises the recovered chiral amine compound.

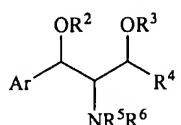
29. (New) The method of Claim 22 further comprising racemizing at least a portion of the second enantiomer in the separated solution by contacting the second enantiomer with a base.

Amdt. dated December 22, 2005

Response to Restriction Requirement and Preliminary
Amendment

30. (New) The method of Claim 29, wherein the enantiomeric mixture of the α -(phenoxy)phenylacetic acid compound used in said step (a) comprises a racemized α -(phenoxy)phenylacetic acid compound.

31. (New) The method of Claim 22, wherein the chiral amine compound is of the formula:



wherein

each of R² and R³ is independently hydrogen or alkyl; or R² and R³ together with atoms to which they are attached to form a heterocyclic ring moiety;

R⁴ is hydrogen or alkyl;

each of R⁵ and R⁶ is independently hydrogen or alkyl, or one of R⁵ or R⁶ is an amine protecting group; and

Ar is aryl.

32. (New) The method of Claim 22, wherein the α -(phenoxy)phenylacetic acid is an enantiomeric mixture of 4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid, said method comprising:

(a) producing a solution comprising an enantiomerically enriched acid-base salt of (-)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid by contacting the enantiomeric mixture of 4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid with less than 0.5 molar equivalent of an enantiomerically enriched (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol in about 4 grams of an alcoholic solvent per gram of (-)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid;

(b) separating the enantiomerically enriched acid-base salt from the solution which is enriched with (+)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid; and

Amdt. dated December 22, 2005

Response to Restriction Requirement and Preliminary
Amendment

(c) removing (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol from the acid-base salt to produce enantiomerically enriched (-)-4-chloro- α -(3-trifluoromethyl-phenoxy)phenylacetic acid.

33. (New) The method of Claim 32, wherein the alcoholic solvent is isopropanol.

34. (New) The method of Claim 33, wherein about 0.47 molar equivalent or less of (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol is used to form the acid-base salt.

35. (New) The method of Claim 34, wherein said step (a) of producing a solution comprising an enantiomerically enriched acid-base salt of (-)-4-chloro- α -(3-trifluoromethyl-phenoxy)phenylacetic acid comprises heating the solution mixture to a temperature at or above a nucleation temperature of the (-)-acid-base salt.

36. (New) The method of Claim 35, wherein said step (b) of separating the enantiomerically enriched acid-base salt is performed at a temperature near or above a saturation temperature of an acid-base salt of the (+)-enantiomer.

37. (New) The method of Claim 33, wherein the enantiomerically enriched (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol comprises at least a portion of (1R,2R)-2-amino-1-(4-nitrophenyl)-1,3-propanediol that is removed from the acid-base salt of said step (c).

38. (New) The method of Claim 33 further comprising racemizing at least a portion of (+)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid obtained in said step (b).

39. (New) The method of Claim 38, wherein the enantiomeric mixture of 4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid comprises at least a portion of (+)-4-chloro- α -(3-trifluoromethylphenoxy)phenylacetic acid that is racemized.

40. (New) An enantiomerically enriched α -(phenoxy)phenylacetate compound made by the method of any one of claims 22 to 39.

Appl. No. 10/656,567

PATENT

Amdt. dated December 22, 2005

Response to Restriction Requirement and Preliminary
Amendment

41. (New) A pharmaceutical composition comprising a pharmaceutically acceptable excipient and a compound of claim 40.